

ARTICLE

Supervised Multidimensional Item Response Theory Modeling of Pediatric Iatrogenic Withdrawal Symptoms

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Item-level data from composite scales can be analyzed with pharmacometric item response theory (IRT) models to improve the quantification of disease severity compared with the use of total composite scores. However, regular IRT models assume unidimensionality, which is violated in the scale measuring iatrogenic withdrawal in children because some items are also affected by pain, undersedation, or delirium. Here, we compare regular IRT modelling of pediatric iatrogenic withdrawal symptom data with two new analysis approaches in which the latent variable is guided towards the condition of interest using numerical withdrawal severity scored by nurses as a “supervising variable.” supervised IRT (sIRT) and supervised multi-dimensional (smIRT) modelling. In this example, in which the items scores are affected by multiple conditions, regular IRT modeling is worse to quantify disease severity than the total composite score, whereas improved performance compared with the composite score is observed for the sIRT and smIRT models.

Study Highlights

WHAT IS THE CURRENT KNOWLEDGE ON THE TOPIC?

☑ The item-level data from composite scales can be analyzed with pharmacometric item response theory (IRT) models, which improves the quantification of the disease severity compared with the total composite score. However, regular IRT models assume unidimensionality, which is not the case for clinical scales of iatrogenic withdrawal because its items can also be affected by pain, undersedation, or delirium.

WHAT QUESTION DID THIS STUDY ADDRESS?

☑ The study analyzes data from a composite scale for monitoring pediatric iatrogenic withdrawal and compares the performance of regular IRT modeling and the following

two novel methods of IRT modeling: supervised IRT (sIRT) and supervised multidimensional IRT (smIRT).

WHAT DOES THIS STUDY ADD TO OUR KNOWLEDGE?

☑ Regular IRT modeling was worse than the total composite score in quantifying iatrogenic withdrawal. Both sIRT and smIRT modeling provided superior quantification of withdrawal compared with the total composite score.

HOW MIGHT THIS CHANGE DRUG DISCOVERY, DEVELOPMENT, AND/OR THERAPEUTICS?

☑ The supervised IRT methods introduced here can improve the analysis of data from composite clinical scales that are not unidimensional.

In the pediatric intensive care unit (ICU), critically ill children often receive opioids and sedatives for prolonged periods of time, which may contribute to drug dependence and iatrogenic withdrawal syndrome (IWS).^{1,2} The importance of careful weaning from these drugs, instead of abrupt discontinuation, is well established. However, even in studies with standardized weaning protocols, IWS is reported in 5–87% of children.^{1,3} Lacking strategies for predicting individualized weaning strategies, weaning is guided by the frequent monitoring for IWS.⁴ However, the lack of specific withdrawal symptoms makes monitoring for IWS difficult because a particular symptomatic profile (e.g., a crying child with tense body parts, anxiety, and hyperalertness)

might be caused by IWS but also by pain, undersedation, or delirium as illustrated in **Figure 1**.^{5–7} In practice, ICU nurses use contextual information to decide on the most likely explanation of the symptoms.^{5,8} For example, IWS might be expected during weaning from opioids.

The Sophia Observational withdrawal Scale (SOS_{withdrawal}) is a validated scale that scores the presence of 15 symptoms related to IWS.^{9,10} The number of symptoms present equals the composite score of this scale, which is used to identify IWS. As such, all symptoms are weighed equally and treated as equally informative of IWS. However, if some symptoms are more informative of IWS than other symptoms, a superior quantification of IWS will be obtained by

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Multidimensional Item Response Theory

Multidimensional Item Response Theory

Jin-Ying Zhang



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